REMARKS

Reconsideration of this application, as amended, is respectfully requested.

The Examiner made the restriction requirement final and withdrew nonelected Claims 8-11 from further consideration. In response, Claims 8-11 are appropriately shown as "withdrawn" in the current amendment. Once again, Applicants reserve the right to file a divisional application directed to the nonelected subject matter of this invention, if desired, in due course.

Applicants gratefully acknowledge that the Examiner kindly indicates in the Office Action Summary that Claims 1-7 and 12-16 are allowed.

In the Detailed Action, the Examiner objects to Claims 18 and 20 as being dependent upon a rejected base claim, but these claims would be allowable if rewritten in independent form including all of the limitations of the base claim and any intervening claims. Base Claims 17 and 19 have been rejected by the Examiner under 35 U.S.C. § 112, second paragraph, as being indefinite due to the phrase "such as." To expedite matters but without comment on the merits of the rejection, Applicants have removed "such as" from Claims 17 and 19 for the better readability thereof. Since this amendment renders the rejection moot and also obviates the objection, it is respectfully asked that the rejection of Claims 17 and 19 as well as the objection to Claims 18 and 20 be withdrawn.

Accordingly, the application is now in condition for an allowance and favorable treatment is respectfully urged.

Respectfully submitted,

WYETH

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APPENDIX

AMENDMENTS TO THE CLAIMS

Please amend the claims as follows:

Claim 1 (Currently amended). A method for synthesis of 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, the process comprising the steps of:

a) reacting 2-(3-Nitro-benzenesulfonylamino)-acetamide with $ClCH_2CONH_2$ ~~N,N-Dimethylformamide~~ in the presence of N,N-Dimethylformamide ~~$ClCH_2CONH_2$~~ and a base to provide 2-[Carbamoylmethyl-(3-nitro-benzenesulfonyl)-amino]acetamide;

b) treating the 2-[Carbamoylmethyl-(3-nitro-benzenesulfonyl)-amino]acetamide product of step a) with a reducing agent to provide 2-[(3-Amino-benzenesulfonyl)-carbamoylmethyl-amino]acetamide;

c) treating the 2-[(3-Amino-benzenesulfonyl)-carbamoylmethyl-amino]acetamide product of step b) with cyanuric chloride to give 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide; and

d) reacting the 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide product of step c) with the disodium salt of 4,4'-diamino-2,2'-biphenyldisulfonic acid.

Claim 2 (Original). The method of Claim 1 wherein the treatment of 2-[(3-Amino-benzene-sulfonyl)-carbamoylmethyl-amino]acetamide with cyanuric chloride to give 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide is conducted at a temperature of from about 20°C to about 25°C.

Claim 3 (Original). The method of Claim 1 wherein the treatment of 2-[(3-Amino-benzene-sulfonyl)-carbamoylmethyl-amino]acetamide with cyanuric chloride to give 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}benzenesulfonyl)-carbamoylmethyl-amino]acetamide is conducted in a reaction medium containing 1-methyl-2-pyrrolidinone and sodium carbonate or sodium bicarbonate.

Claim 4 (Original). The method of Claim 1 further comprising the step of recrystallizing the 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzene-sulfonyl)-carbamoylmethyl-amino]acetamide product of step c) from a mixture of 1-methyl-2-pyrrolidinone and water prior to completing the reaction of step d).

Claim 5 (Original). The method of Claim 1 wherein step d) is conducted at a temperature of from about 15°C to about 90°C.

Claim 6 (Original). The method of Claim 5 wherein step d) is conducted at a temperature of from about 60°C to about 75°C.

Claim 7 (Original). The method of Claim 1 wherein step d) is conducted in a medium comprising dimethyl sulfoxide.

Claim 8 (Withdrawn).

Claim 9 (Withdrawn).

Claim 10 (Withdrawn).

Claim 11 (Withdrawn).

Claim 12 (Original). A process for purifying 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide

obtained by treating 2-[(3-amino-benzenesulfonyl)-carbamoylmethyl-amino]acetamide with cyanuric chloride, which comprises dissolving 2-[(4-{4-[4-(bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide in a volume of water and 1-methyl-2-pyrrolidinone, followed by addition of excess water to precipitate a more purified amount of 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide.

Claim 13 (Currently amended). The ~~A~~ process according to Claim ~~claim~~ 12 wherein the ratio of water:1-methyl-2-pyrrolidinone into which the amount of 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]-acetamide is dissolved is from about 1:1 by weight.

Claim 14 (Currently amended). The ~~A~~ process according to Claim ~~claim~~ 12 wherein precipitation of the desired 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide product is carried out by adding additional water to create a water:1-methyl-2-pyrrolidinone ratio of up to about 6:1 (wt:wt)~~;~~.

Claim 15 (Currently amended). The ~~A~~ process according to Claim ~~claim~~ 13 wherein precipitation of the desired 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide product is carried out by adding additional water to create a water:1-methyl-2-pyrrolidinone ratio of up to about 6:1 (wt:wt)~~;~~.

Claim 16 (Currently amended). The ~~A~~ process according to Claim ~~claim~~ 14 wherein precipitation of the desired 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide product is carried out by adding additional water to create a water:1-methyl-2-pyrrolidinone ratio from about 3:1 to about 5:1 (wt:wt).

Claim 17 (Currently amended). A process for the increasing the purity of 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, disodium salt which comprises dissolving impure 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, disodium salt in volume of acetonitrile-water having a mixture ratio of from about 0.75:2 to about 1.5:2 by volume at an elevated temperature~~, such as of~~ from about 30°C to about 70°C, followed by addition of additional acetonitrile until crystallization of the desired compound is achieved.

Claim 18 (Currently amended). ~~The A~~ process according to Claim 17 wherein the elevated temperature is from about ~~65~~ 60°C - about 70°C and after the addition of additional acetonitrile the mixture is cooled to about 49°C - about 51°C.

Claim 19 (Currently amended). ~~The A~~ process according to ~~Claim claim~~ 1 in which the 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, disodium salt prepared is purified by dissolving impure 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, disodium salt in volume of acetonitrile-water having a mixture ratio of from about 0.75:2 to about 1.5:2 by volume at an elevated temperature~~, such as of~~ from about 30°C to about 70°C, followed by addition of additional acetonitrile until crystallization of the desired compound is achieved.

Claim 20 (Currently amended). ~~The A~~ process according to ~~Claim claim~~ 19 in which the 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, disodium salt prepared is purified by dissolving impure 4',4-bis-{4,6-bis-[3-(bis-carbamoyl-methyl-1-sulfamoyl)-phenylamino]-[1,3,5]triazin-2-ylamino}-biphenyl-2,2'-disulfonic acid, disodium salt in volume of acetonitrile-water having a mixture ratio of from about 0.75:2 to about 1.5:2 by volume at an elevated temperature of from about ~~65~~ 60°C to about 70°C, followed by addition of additional acetonitrile and cooling of the mixture to a temperature of from about 49°C to about 51°C until crystallization of the desired compound is achieved.

Please add the following claim:

Claim 21 (New). The process according to Claim 16 wherein precipitation of the desired 2-[(4-{4-[4-(Bis-carbamoylmethyl-sulfamoyl)-benzyl]-6-chloro-[1,3,5]triazin-2-ylmethyl}-benzenesulfonyl)-carbamoylmethyl-amino]acetamide product is carried out by adding additional water to create a water:1-methyl-2-pyrrolidinone ratio of about 4:1 (wt:wt).